

WHAT IS CLAIMED IS:

1. An isolated variant allele of a human kappa opioid receptor gene, comprising a DNA  
2 sequence having at least one variation in SEQ ID NO:1, wherein said variation  
3 comprises C852T, C948T, or C1008T, or any combination thereof.
1. The isolated variant allele of Claim 1, detectably labeled.
1. The isolated variant allele of Claim 2, wherein said detectable label comprises a  
2 radioactive element, a chemical which fluoresces, or an enzyme.
1. An isolated nucleic acid molecule selectively hybridizable to the isolated variant allele  
2 of Claim 1.
1. The isolated nucleic acid molecule of Claim 4, detectably labeled.
1. The isolated nucleic acid molecule of Claim 5, wherein said detectable label comprises a  
2 radioactive element, a chemical that fluoresces, or an enzyme.
1. A cloning vector comprising an isolated variant allele of a human kappa opioid receptor  
2 gene and an origin of replication, wherein said variant allele comprises a DNA sequence  
3 having at least one variation in SEQ ID NO:1, wherein said variation comprises C852T,  
4 C948T, or C1008T, or any combination thereof.
1. A cloning vector comprising an origin of replication and an isolated nucleic acid  
2 molecule selectively hybridizable to an isolated variant allele of a human kappa opioid  
3 receptor gene, wherein said variant allele comprises a DNA sequence having at least one  
4 variation in SEQ ID NO:1, wherein said at least one variation comprises C852T, C948T,  
5 or C1008T, or any combination thereof.
1. The cloning vector of either of Claims 7 or 8, wherein said cloning vector comprises of  
2 *E. coli*, bacteriophages, plasmids, or pUC plasmid derivatives.

1 10. The cloning vector of Claim 9, wherein bacteriophages further comprise lambda  
2 derivatives, plasmids further comprise pBR322 derivatives, and pUC plasmid derivatives  
3 further comprise pGEX vectors, or pmal-c, pFLAG.

1 11. An expression vector comprising an isolated variant allele of a human kappa opioid  
2 receptor gene comprising a DNA sequence having a variation in SEQ ID NO:1, wherein  
3 said variation comprises C852T, C948T, or C1008T, or any combination thereof.

1 12. An expression vector comprising an isolated nucleic acid molecule selectively  
2 hybridizable to an isolated variant allele of a human kappa opioid receptor gene, wherein  
3 said isolated nucleic acid molecule is operatively associated with a promoter, and said  
4 variant allele comprises a DNA sequence having at least one variation in SEQ ID NO:1,  
5 wherein said at least one variation comprises C852T, C948T, or C1008T, or any  
6 combination thereof.

1 13. The expression vector of either of Claims 11 or 12, wherein said promoter comprises  
2 immediate early promoters of hCMV, early promoters of SV40, early promoters of  
3 adenovirus, early promoters of vaccinia, early promoters of polyoma, late promoters of  
4 SV40, late promoters of adenovirus, late promoters of vaccinia, late promoters of  
5 polyoma, the *lac* the *trp* system, the *TAC* system, the *TRC* system, the major operator  
6 and promoter regions of phage lambda, control regions of fd coat protein, 3-  
7 phosphoglycerate kinase promoter, acid phosphatase promoter, or promoters of yeast  $\alpha$   
8 mating factor.

1 14. A unicellular host transformed or transfected with an expression vector comprising an  
2 isolated variant allele of a human kappa opioid receptor gene operatively associated with  
3 a promoter, wherein said variant allele comprises a DNA sequence having at least one  
4 variation in SEQ ID NO:1, wherein said at least one variation comprises C852T, C948T,  
5 or C1008T, or any combination thereof.

1 15. A unicellular host transformed with an expression vector comprising an isolated nucleic  
2 acid molecule selectively hybridizable to an isolated variant allele of a human kappa  
3 opioid receptor gene, wherein said isolated nucleic acid molecule is operatively

associated with a promoter, and said variant allele comprises a DNA sequence having at least one variation in SEQ ID NO:1, wherein said at least one variation comprises C852T, C948T, or C1008T, or any combination thereof.

16. The unicellular host of either of Claims 14 or 15, wherein said host comprises *E. coli*, *Pseudomonas*, *Bacillus*, *Streptomyces*, yeast, CHO, R1.1, B-W, L-M, COS1, COS7, BSC1, BSC40, BMT10 or SF9 cells.

17. A method for determining a susceptibility in a subject to at least one physiological response, condition or disease related to the endogenous opioid system, nociception, neurotransmitter release endogenous opioid system, learning, memory and cognition; cocaine, amphetamine and other stimulants self-administration; behavioral sensitization to cocaine, opiates, alcohol and tobacco; opiate, amphetamine and alcohol withdrawal, physical dependence and tolerance; neuroendocrine function, reproductive function, prolactin regulation, stress responsivity; physiology and pathology of mood and affect; immune function, and gastrointestinal function, comprising the steps of:

- removing a bodily sample from said subject, wherein said sample comprises a first and second allele comprising a human kappa opioid receptor gene;
- determining whether said human kappa opioid receptor gene of said first allele comprises a DNA sequence having at least one variation in SEQ ID NO:1, wherein said variation comprises C852T, C948T, or C1008T, or any combination thereof, such that the presence of said at least one variation in said human kappa opioid receptor gene of said first allele is expected to be indicative of the subject's susceptibility to at least one said physiological response, condition or disease relative to the susceptibility to said at least one said physiological response, condition or disease in a standard.

18. The method for determining a susceptibility to at least one addictive disease of Claim 17, further comprising the step of determining whether said human kappa opioid receptor gene of said second allele comprises a DNA sequence having at least one variation in SEQ ID NO:1, wherein said variation comprises C852T, C948T, or C1008T, or any combination thereof, such that the presence of said at least one variation in said human kappa opioid receptor gene of said second allele is expected to be indicative of the

7 subject's susceptibility to said at least one physiological response, condition or disease  
8 relative to the susceptibility to said at least one physiological response, condition or  
9 disease in said standard.

1 19. The method of either of Claim 19 wherein said at least one addictive disease comprises  
2 opioid addiction; cocaine addiction or addiction to other psychostimulants; nicotine  
3 addiction; barbituate or sedative hypnotic addiction; anxiolytic addiction; or  
4 alcohol addiction.

1 20. A method for determining a susceptibility to pain in a subject relative to a susceptibility  
2 of pain in a standard, wherein the method comprises the steps of:  
3 a) removing a bodily sample from said subject, wherein said sample comprises a  
4 first and second allele comprising a human kappa opioid receptor gene;  
5 b) determining whether said human kappa opioid receptor gene of said first allele  
6 comprises a DNA sequence having at least one variation in SEQ ID NO:1,  
7 wherein said variation comprises C852T, C948T, or C1008T, or any  
8 combination thereof, such that the presence of said at least one variation in said  
9 human kappa opioid receptor gene of said first allele is expected to be indicative  
10 of susceptibility to pain in said subject relative to susceptibility to pain in said  
11 standard, wherein said first allele of said standard comprises a human kappa  
12 opioid receptor gene comprising a DNA sequence of SEQ ID NO:1.

13 21. The method of Claim 20 for determining a susceptibility to pain in a subject, further  
1 comprising the step of determining whether said second allele of said bodily sample  
2 comprises a human kappa opioid receptor gene comprising a DNA sequence having at  
3 least one variation in SEQ ID NO:1, wherein said variation comprises C852T, C948T, or  
4 C1008T, or any combination thereof, such that the presence of said at least one variation  
5 in said second allele is expected to be indicative of susceptibility to pain in said subject  
6 relative to susceptibility of pain in said standard, wherein said second allele of said  
7 standard comprises a human kappa opioid receptor gene comprising a DNA sequence of  
8 SEQ ID NO:1.

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1 22. A method for determining a therapeutically effective amount of pain reliever to  
2 administer to a subject in order to induce analgesia in said subject relative to a  
3 therapeutically effective amount of pain reliever to administer to a standard in order to  
4 induce analgesia in said standard, wherein the method comprises determining a  
5 susceptibility to pain in said subject relative to susceptibility to pain in said standard,  
6 wherein susceptibility to pain in said subject is expected to be indicative of said  
7 therapeutically effective amount of pain reliever to administer to said subject to induce  
8 analgesia in said subject relative to said therapeutically effective amount of pain reliever  
9 to administer to said standard to induce analgesia in said standard.

1 23. The method of Claim 22 for determining a therapeutically effective amount of pain  
2 reliever to administer to said subject, wherein determining susceptibility to pain in said  
3 subject comprises the steps of:  
4 a) removing a bodily sample from said subject, wherein said sample comprises a  
5 first and second allele comprising a human kappa opioid receptor gene; and  
6 b) determining whether said first allele comprises a human kappa opioid receptor  
7 gene comprising a DNA sequence having at least one variation in SEQ ID NO:1,  
8 wherein said at least one variation comprises C852T, C948T, or C1008T, or any  
9 combination thereof, wherein the presence of said at least one variation in said  
10 human kappa opioid receptor gene of said first allele is expected to be indicative  
11 of the subject's susceptibility to pain relative to said to susceptibility of pain in  
12 said standard, wherein said first allele of said standard comprises a human kappa  
13 opioid receptor gene comprising a DNA sequence of SEQ ID NO:1, such that  
14 said therapeutically effective amount of pain reliever to administer to the subject  
15 in order to induce analgesia is related to said susceptibility to pain in said subject  
16 relative to susceptibility to pain in said standard.

1 24. The method of Claim 23, wherein determining susceptibility to pain in said subject  
2 relative to susceptibility to pain in said standard further comprises the step of  
3 determining whether said second allele of said bodily sample from said subject  
4 comprises a human kappa opioid receptor gene comprising a DNA sequence having at  
5 least one variation in SEQ ID NO:1, wherein said at least one variation comprises  
6 C852T, C948T, or C1008T, or any combination thereof, such that the presence of said at

7 least one variation in said second allele is expected to be indicative of susceptibility to  
8 pain in said subject relative to susceptibility to pain in said standard, wherein said second  
9 allele of said standard comprises a human kappa opioid receptor gene comprising a DNA  
10 sequence of SEQ ID NO:1, and the therapeutically effective amount of pain reliever to  
11 administer to said subject to induce analgesia in said subject is related to the presence of  
12 said at least one variation in said human kappa opioid receptor gene of said second allele  
13 of said bodily sample from said subject.

1 25. A method for determining a therapeutically effective amount of therapeutic agent to  
2 administer to a subject suffering from at least one addictive disease to treat the at least  
3 one addictive disease in said subject relative to a therapeutically effective amount of  
4 therapeutic agent to administer to a standard suffering from the at least one addictive  
5 disease to treat the at least one addictive disease in said standard, wherein the method  
6 comprises the steps of:  
7 a) removing a bodily sample from said subject, wherein said sample comprises a  
8 first and second allele comprising a human kappa opioid receptor gene; and  
9 b) determining whether said first allele comprises a human kappa opioid receptor  
10 gene comprising a DNA sequence having at least one variation in SEQ ID NO:1,  
11 wherein said variation comprises C852T, C948T, or C1008T, or any  
12 combination thereof,  
13 wherein the presence of said at least one variation in said human kappa opioid receptor  
14 gene of said first allele is expected to be indicative of the therapeutically effective  
15 amount of said therapeutic agent to administer to the subject to treat said at least one  
16 addictive disease in said subject relative to said therapeutically effective amount of said  
17 therapeutic agent to administer to said standard to treat said at least one addictive disease  
18 in said standard, wherein said first allele of said standard comprises a human kappa  
19 opioid receptor gene comprising a DNA sequence of SEQ ID NO:1.

1 26. The method of Claim 25 for determining a therapeutically effective amount of  
2 therapeutic agent to administer to a subject suffering from said at least one addictive  
3 disease to treat said at least one addictive disease, relative to said therapeutically  
4 effective amount of said therapeutic agent administered to said standard suffering from  
5 said at least one addictive disease to treat said at least one addictive disease in said

6 standard, further comprising the step of determining whether said second allele of said  
7 bodily sample from said subject comprises a human kappa opioid receptor gene  
8 comprising a DNA sequence having at least one variation in SEQ ID NO:1, wherein said  
9 variation comprises C852T, C948T, or C1008T, or any combination thereof, such that  
10 the presence of said at least one variation in said second allele related to said  
11 therapeutically effective amount of said therapeutic agent administered to said subject to  
12 treat said at least one addictive disease in said subject relative to said therapeutically  
13 effective amount of said therapeutic agent to administer to said standard to treat said at  
14 least one addictive disease in said standard, wherein said second allele of said standard  
15 comprises a human kappa opioid receptor gene comprising a DNA sequence of SEQ ID  
16 NO:1.

1 27. The method of either of Claims 25 or 26, wherein said at least one addictive disease  
2 comprises opioid addiction; cocaine addiction or addiction to other psychostimulants;  
3 nicotine addiction; barbiturate or sedative hypnotic addiction; anxiolytic addiction; or  
4 alcohol addiction.

5 28. A commercial test kit for determining the presence of at least one variation in a  
6 human kappa opioid receptor gene of an allele in a bodily sample taken from a subject,  
7 wherein the commercial test kit comprises:  
8 a) PCR oligonucleotide primers suitable for detection of an allele comprising a  
9 human kappa opioid receptor gene comprising a DNA sequence having at least  
one variation in SEQ ID NO:1 comprising C852T, C948T, or C1008T, or any  
combination thereof;  
b) other reagents; and  
c) directions for use of the kit.

1 29. A nucleic acid as set forth in SEQ ID No:1.